- Fig. 2 shows the synthetic scheme for synthesis of an intermediate in the preparation of the derivative of Fig. 1;

- Fig. 3 shows HPLC chromatograms over a period of time in a hydrolysis test of a cyclosporin derivative;

B1 Conclude

- Fig. 4 is a curve showing the variation with time of the concentration of the products in the same hydrolysis test; and

- Fig. 5 is a curve showing the kinetics of inhibition, by a cyclosporin derivative, of cis-trans isomerase activity in Cyclophilin A from calf thymus.

DETAILED DESCRIPTION OF THE INVENTION

Please replace the paragraph beginning at page 4, line 17, with the following rewritten paragraph:

B2 -- The properties of the cyclosporin derivatives of the present invention, the advantages offered by them, and the detailed method of preparation of these derivatives will be illustrated using the specific examples below, and with the help of the drawings. --

Page 4, please delete lines 22 - 32.

Page 5, please delete lines 1 - 3.

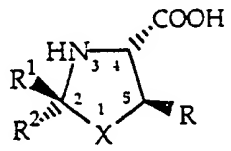
IN THE CLAIMS

Please substitute the following amended claims for corresponding claims previously presented. A copy of the amended claims showing current revisions is attached.

Kindly amend the following claims.

B3

1. (Amended) A cyclosporin derivative in which the peptide chain comprises at least one residue of a non-natural amino acid of formula I:



(I)

in which

X denotes an oxygen or a sulfur;

R denotes a hydrogen, or an alkyl group having between 1 and 6 carbon atoms;

R_1 and R_2 denote, independently of each other, a hydrogen, an alkyl group, having between 1 and 6 carbons, which may be straight-chain or branched-chain, substituted or non-substituted, an alkylene group having between 1 and 6 carbon atoms, a substituted or non-substituted aryl group, a substituted or non-substituted heteroaryl group, a residue of a water-soluble polymer, possibly bound to a spacer group.

2. (Amended) The derivative according to Claim 1, wherein in the amino acid of formula I, R denotes a hydrogen or a methyl group.

B4

3. (Twice Amended) The derivative according to Claim 1, wherein it is derived from a cyclosporin in which the peptide chain contains at least one amino acid, chosen from serine, threonine and cysteine, in D or L configuration.